

خير الناس أنفعهم للناس

تجميعات

ALKALOIDS

BY/A.H.A

لا تنسونا من صالح دعائكم
بظهر الغيب

Tyrosine $\xrightarrow{\text{derived from}}$ I phenylalkylamine alkaloids			II Pyridine alkaloids			
① Ephedra alkaloids	② Khat alkaloids	③ Peyote alkaloids	i) only Trigonelline	ii) Tetrahydro pyridine alk.	iii) + other N base	iv) Pyridone alkaloids
<chem>CC(N)Cc1ccccc1</chem> (L-ephedrine) \rightarrow ephedra herb \rightarrow volatile, sol. in H_2O \rightarrow so ppt & mayer's (w/ Ca^{2+}) NB! - ethyl nitrate in extract? as ephedrine in aged chloroform, test sol. by benzene - up to 10% & crystalline and forming phosgene toxic gas.	<chem>CC(N)Cc1ccccc1</chem> (Cathinone) (Cathine) \rightarrow Khat leaves	<chem>CC(N)Cc1ccccc1</chem> (mescaline) \rightarrow Peyote \rightarrow Sol. in H_2O organic solvent	<chem>CC1Cc2ccccc2N1</chem> (Trigonelline) \rightarrow Foerugreek seed	<chem>CC1Cc2ccccc2N1</chem> (Guaicoline) \rightarrow (Guaicoline) \rightarrow (Atracoline) \rightarrow Atracine nut	<chem>CC1Cc2ccccc2N1</chem> (nicotine) (nicotinic) (nicotamine) \rightarrow Volatile & liquid \rightarrow (nicotinic) (nicotelline) (nicotelline) \rightarrow Tobacco leaves	<chem>CC1Cc2ccccc2N1</chem> (Ricinine) \rightarrow Castor seed
① antiasthmatic	① CNS stimulant (Hallucinogenic) \rightarrow Cause Habitual and addictive	① Hallucinogenic & psychomimetics \rightarrow not cause Habitual and addictive	① Hypoglycemic	① Anthelmintic (2) CNS stimulant (dose)	① Insecticide (2) Toxin in dose (pesticide)	① poisonous (CN) (Toxic alkaloid)
② Nasal decongestant (2) Sympathetic stimulant	② Chen's Test $+ CuSO_4 + NaOH \rightarrow$ Violet color either \rightarrow purple color in ether layer		② Seeds deffing Mark Free base \rightarrow Meq & Total alk.	② + K Ferri cyanide \rightarrow Blue ② + K Ferri cyanide \rightarrow green	② + PMAB \rightarrow blood color	② + $KMnO_4 \rightarrow$ decoloring
Synthesis of ephedrine <chem>CC(=O)O + CH_3-CH_2-NH_2 \xrightarrow{H_2SO_4} CC(=O)O-CH_2-CH_2-NH_2</chem>	<chem>CC(=O)O + CH_3-CH_2-NH_2 \xrightarrow{H_2SO_4} CC(=O)O-CH_2-CH_2-NH_2</chem>	<chem>CC(=O)O + CH_3-CH_2-NH_2 \xrightarrow{H_2SO_4} CC(=O)O-CH_2-CH_2-NH_2</chem>		② + K Ferri cyanide \rightarrow Blue ② + K Ferri cyanide \rightarrow green	② + PMAB \rightarrow blood color	② + $KMnO_4 \rightarrow$ decoloring
④ Colchicum alkaloids	⑤ Capsicum alkaloids	⑥ Tropane alkaloids	V Piperidine alkaloids			
<chem>CC1Cc2ccccc2N1</chem> (Colchicine) \rightarrow w. base \rightarrow Colored alkaloids (yellow) \rightarrow Sol. in H_2O \rightarrow Calchicine + CH_3	<chem>CC1Cc2ccccc2N1</chem> (Capsaicin) \rightarrow phenolic amide alkaloids \rightarrow pungent alkaloids (Pungency destroyed by $KMnO_4$) \rightarrow in sol. in H_2O	<chem>CC1Cc2ccccc2N1</chem> (Tropane) \rightarrow non volatile liquid \rightarrow Sol. in H_2O , CH_2Cl_2 \rightarrow insol. in ether \rightarrow tropane alkaloids	a) Pepper alkaloids <chem>CC1Cc2ccccc2N1</chem> (Piperine) \rightarrow Hemlock Fruit (Gonium) \rightarrow Volatile, liquid b) Conium alkaloids <chem>CC1Cc2ccccc2N1</chem> (Coniine) \rightarrow Hemlock Fruit (Gonium) \rightarrow Volatile, liquid c) Lobelia alkaloids <chem>CC1Cc2ccccc2N1</chem> (Lobeline) \rightarrow Lobelia herb d) Pomegranate alkaloids <chem>CC1Cc2ccccc2N1</chem> (Punicic acid) \rightarrow Pomegranate			
① Treatment of G-out	① Rubefacient (counter irritant)	① Miotic & choleragic act	Synthesis of lobeline			
② plant hormone	② anti rheumatic	② UL of glaucoma	<chem>CC1Cc2ccccc2N1</chem> (Lobeline) \rightarrow Lobelia herb (Lobeline) \rightarrow Lobelia herb (Lobeline) \rightarrow Lobelia herb			
Colchicine + Fecl ₃ \rightarrow Red color Colchicine + Fecl ₃ \rightarrow olive green color	+ Fecl ₃ \rightarrow green color	Helch's Test $+ H_2O_2 + K_2Cr_2O_7 \rightarrow$ violet color yellowish brown \rightarrow violet color (in organic layer)	(Benzoyl) \rightarrow <chem>CC1Cc2ccccc2N1</chem> (Lobeline) \rightarrow Lobelia herb (Lobeline) \rightarrow Lobelia herb			
			(Benzoyl) \rightarrow <chem>CC1Cc2ccccc2N1</chem> (Lobeline) \rightarrow Lobelia herb (Lobeline) \rightarrow Lobelia herb			



pyrrolidine + piperidine) \leftarrow **IV Tropicane** alkaloids $\xrightarrow{\text{derived from}}$ L-ornithine

① official reaction of Rane's compound
② ester alkaloids

V Quinine alkaloids

A Solanaceous alkaloids

ester

B CoCa alkaloids

A Cinchona alkaloids

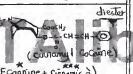
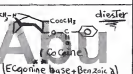
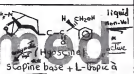
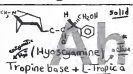
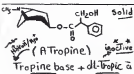
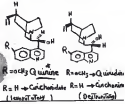
i) Atropine

ii) Hyoscyamine

iii) Hyoscyne

i) Cocaine

ii) Cinnamyl Cocaine



Solanaceous leaves

CaCa leaves

Cinchona bark

- ① Anticholinergic effect = Mydriatic = Parasympatholytic
- ② Antispasmodic \rightarrow smooth muscle relaxant
- ③ \downarrow secretion
- ④ Antidote for organophosphorus insecticides

- ① Sedative
- ② CNS depressant

- ① Local anesthetic (dentistry)
- ② CNS stimulant

- ① + KMnO₄ \rightarrow decolorization
- ② + KMnO₄ \rightarrow violet

- ① Antimalarial \rightarrow Quinine
- ② Anticholinergic \rightarrow Quinidine

Tests

① Vital's marin's Test (+ve for all)

② Geyard's Test

③ + KMnO₄ \rightarrow violet

④ + KMnO₄ \rightarrow decolorization

⑤ Fluorescence Test

alk + Fuming HNO₃ \rightarrow violet color

alk + H₂O₂/alkaloid \rightarrow Red color (Hyoscyamine)

alk + KMnO₄ \rightarrow violet

alk + KMnO₄ \rightarrow decolorization

alk + KMnO₄ \rightarrow decolorization

alk + KMnO₄ \rightarrow decolorization

③ Shaw's Test: \rightarrow green color

④ PDAB Test: \rightarrow Red color

⑤ + KMnO₄ \rightarrow violet

⑥ + KMnO₄ \rightarrow decolorization

⑦ + KMnO₄ \rightarrow decolorization

⑧ + KMnO₄ \rightarrow decolorization

① Strong base

② Hyoscyamine

③ Weak base

④ + KMnO₄ \rightarrow violet

⑤ + KMnO₄ \rightarrow decolorization

⑥ + KMnO₄ \rightarrow decolorization

Quen. deterng of i) ii) iii)

Isolatn

Synthetic of Atropine

Synthetic of Cocaine

Semisynthesis of Cocaine

Isolatn

① Total alkaloids by acid base titration

② R/o of each alkaloids

③ Total alkaloids by acid base titration

④ Total alkaloids by acid base titration

⑤ Total alkaloids by acid base titration

⑥ Total alkaloids by acid base titration

⑦ Total alkaloids by acid base titration

⑧ Total alkaloids by acid base titration

⑨ Total alkaloids by acid base titration

⑩ Total alkaloids by acid base titration

⑪ Total alkaloids by acid base titration

⑫ Total alkaloids by acid base titration

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㋗ Total alkaloids by acid base titration

㋘ Total alkaloids by acid base titration

㋙ Total alkaloids by acid base titration

㋚ Total alkaloids by acid base titration

㋛ Total alkaloids by acid base titration

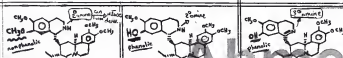
3

Isaquinoline alkaloids

Benzyl isoquinoline GP → Papaverine
Phenyl isoquinoline GP → Narcotine
Phenanthrene (Morphine) GP → Morphine (Morphine)

A Ipecacuanha alkaloids

① Emetine ← Methyl ② Cephaline ← Reduct ③ Psychotrine



Insol. in H₂O (Bulky quinuclidine)
① Insol. in NaOH ① Sol. in NaOH
② Sol. in ether ② Insol. in ether

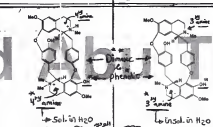
Uses ① expectorant (Emetine drug)
② Antiamoebic of Feat (III of amoebic dysentery) → (emetine HCl)
③ emetic ④ anti viral & anti tumor

Test ① alk. soln + Ca. Hypochlorite → orange color
② Froehde's reagent test → Emetine and cephaline → dirty green color
→ Psychotrine → Pale green color

اللهم دير لنا فينا لا تحسرنا لنديرنا آمين

B Curate alkaloids

① d-Tubocurarine ② Curine



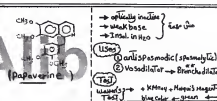
① Sol. in H₂O ① d-Form (active)
② Sol. in H₂O ② Bolus in Curate extract

USES ① skeletal muscle relaxant ① Tubocurarine (active)
② antidote for strychnine poisoning & tetanus poisoning
Test + FeCl₃ → Faint green color → green color

Isolate ① eq. extract + NH₄OH → (F) (Tubocurarine)
② eq. extract + HCl → (F) (Curine)
(Tubocurarine HCl) (Curine HCl) Filter

C opium alkaloids

① Papaverine



① optically inactive
② weak base
③ Insol. in H₂O

Uses ① antispasmodic (spasmodic)
② vasodilator → Branchial dilator (Papaverine)
Test ① Murexide test → blue color
② Murexide test → blue color

② Morphine ③ codeine
① Insol. in H₂O, ether, cyclo
① phenolic & alcoholic
② Insol. in H₂O, ether, cyclo
① non phenolic & alcoholic
② Sol. in NH₄OH (due to formation of cationic amino complex)

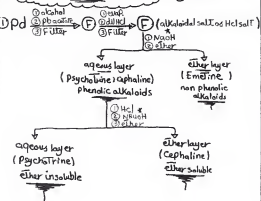
Uses ① Narcotic analgesic
② Cancer pain, Traumatic pain
③ Ill. of Heroin addiction ② anti-tussive

INB/ → Morphine → Heroin (diacetylmorphine)
Morphine → Heroin (diacetylmorphine)

Structure activity Relationship (SAR) of Morphine

↑ activity by: ① Aromatic ring
② Reductive methylation, oxidation
③ elimination of OH at C-3
④ ~ ~ ~ ~ ~ CH₃ at C-4
⑤ ~ ~ ~ ~ ~ CH₃ at C-5
⑥ Replacement of N-CH₃ by N-CH₂CH₃

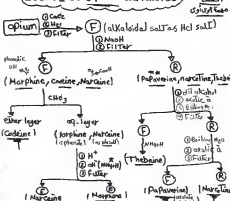
Isolation of Ipeca alkaloids



Notes on opium

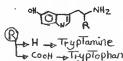
Natural opoids ① Morphine ② Codeine ③ Thebaine
From its use in Form following
Semi-synthetic opoids ① Nalorphine ② Naloxone ③ Buprenorphine ④ Etorphine

Isolation of opium alkaloids



derivate \rightarrow Tryptophan

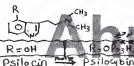
① Serotonin alkaloids



51 Mammalian brain

Propyl
① SKIN IRRITANT $\xrightarrow{\text{H}_2\text{O}}$ Histamine release
② ↑ conc of 5-OH indole acetic acid
↑ histamine Tumor → indicates (Secretion metabolic)

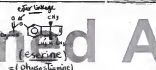
② Psilocin & Psilocybin alkaloids



54 Psilocybe Fungus

① Insol. in H_2O	① Sol. in H_2O
② Sol. in alcohol	② Insol. in $CHCl_3$

(3) Calabar bean colloids

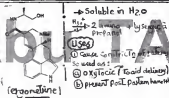


in Calabar bear

① Insol. in H_2O (3M) غير محلول في الماء (3M)
 ② Sol. in $CHCl_3$ (soluble) محلول في $CHCl_3$ (قابل للذوبان)

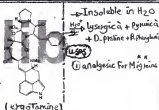
④ Ergot alkaloids

① Ergometrine



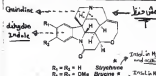
↳ It has certain advantages over the water insoluble ergot alkaloids:
as it does not produce nausea & vomiting
↳ has oxytocic effect

② Ergotamine



Source → Ergot Fungus

⑤ *Nux vomica* alkaloids
(strychnos alkaloids)



in \rightarrow Nux vomica seed

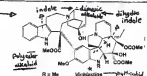
① CNS Stimulant and Tonic. } Stimulant
② antidote for Barbiturate poisoning }
③ Rodenticide ④ Toxic }
① alcohol and oil solvent }
② additive agent to lubricants }
③ less Toxic }

٢٤) من افعال الواح الجاهزة من ممكن

① Styrene + Mandelin's \rightarrow Violet

② Brucine + $\text{SnCl}_2 \rightarrow$ violet

⑥ Vinca alkaloids
(Cathartics alkaloids)



un- Miofa herb

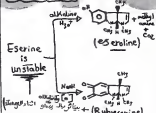
① Tt. of Hodgkin's disease and Carcinoma } vincristine
② anti Cancer (cytotoxic effect)

① Tt. of Child hood leukemia } vincristine
② anti Cancer (cytotoxic effect)

① vinblastine + Vanillin / HCl → Pink color

② Van Wijk's Test (For both) \rightarrow H_2O
 \rightarrow Reddish brown color.

NB on exercise

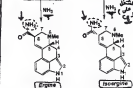


Semi-synthesis deriv of Vinc. alkaloids

Lysergic Acid derivative

eg ergotamine and ergometrine
 ↓
 ↑ Active metabolite
 ↓
 -ine

(-)-Lysergic acid (+)-isolysergic acid



Isolyseric acid derivative

⑧ ergatamine and ergometrine
↓
active Dextro rotating
↓
inine

① Active ergot alkaloids (Fluorescent cpds) $\xrightarrow{\text{light/UV light}}$ Inactive lumi derivatives (non fluorescent cpds)

عقار العنبرية (المطهر)
② LSD (lysergic acid diethylamide) $\xrightarrow{13}$ synthetic cpd (↑ Hallucinogenic effect)

Test

alk. Solg + Van-Wrk's Reagent \longrightarrow deep Blue Color
($\text{POAB} + 15.1. \text{H}_2\text{SO}_4 + \text{Fecl}_3$)

Estimate of ergot alkaloids

by

- ① Colorimetrically
- ② Fluorimetrically

VIII. Purine alkaloids.

① Caffeine	② Theobromine	③ Theophylline
<chem>CN1C=NC2=C1C(=O)N(C)C(=O)N2C</chem>	<chem>CN1C=NC2=C1C(=O)NC(=O)N2C</chem>	<chem>CN1C=NC2=C1C(=O)N(C)C2=NC</chem>
Tea Leaves	Coffee Beans	Cocoa Beans
① weak base → Form salt only w strong acids ② give no ppt w Mayer's Test	① give ppt w Mayer's Test ② slightly sol. in H ₂ O	① give +ve w Murexide Test ② Sol. in H ₂ O and NH ₄ OH
③ Sol. in H ₂ O and Benzene	③ Insol. in Benzene and NH ₄ OH	③ Insol. in Benzene
① CNS Stimulant ② Headache	① Diuretic	① smooth muscle relaxant → Bronchodilator
① Murexide Test (+ve for all) ② Tannic Acid Test (+ve for ① & ③) ③ FeSO ₄ Test (+ve for ②)	① alk. + C.HCl + KClO ₄ → Red color ② alk. sol. + Tannic acid → white ppt (soluble in 33% aq. sol) ③ alk. sol. + C.HCl + Br ₂ /H ₂ O + FeSO ₄ + NH ₃ → Blue color	① alk. + C.HCl + KClO ₄ → Red color ② alk. sol. + Tannic acid → white ppt (soluble in 33% aq. sol) ③ alk. sol. + C.HCl + Br ₂ /H ₂ O + FeSO ₄ + NH ₃ → Blue color

Separation of Purine Alkaloids

mix + Benzene → (F) Caffeine
 (F) Caffeine + NH₄OH → (D) Theophylline
 (D) Theophylline + NH₄OH → (R) Theobromine

IX. Steroidal alkaloids

① Solanine	② Solasoline (Solasodine)
<chem>CN1[C@H]2CC[C@@H]3[C@H]4CC[C@@H]5[C@@]4(CC[C@H]3[C@H]2CC=C4[C@@]5(CC[C@@H](C4)O)C)C</chem>	<chem>CN1[C@H]2CC[C@@H]3[C@H]4CC[C@@H]5[C@@]4(CC[C@H]3[C@H]2CC=C4[C@@]5(CC[C@@H](C4)O)C)C</chem>
(Solanine)	(Solasoline)
→ insol. in H ₂ O → Sol. in aq. sol; Ethyl alcohol	→ Sol. in H ₂ O → Jelly on Cooling
Solanidine + D-glucose + D-galactose + L-Rhamnose	Solasodine + D-glucose + D-galactose + L-Rhamnose
① + Marquis' reagent → yellow color ② + Mandelin's reagent → orange color	① + Marquis' reagent → Red color ② + SbCl ₃ → Red color

Starting material for synthesis of Steroidal drugs (aglycone part) is use

XI. Carboline alkaloids

① Yohimbin alkaloids	② Rauwolfia alkaloids
<chem>O=C1C=CC2=C1C(=O)N(C)C2</chem>	<chem>O=C1C=CC2=C1C(=O)N(C)C2</chem>
(Yohimbine)	(Rauwolfia Root)
in Yohimbin bark	in Rauwolfia Root
Uses → Aphrodisiac	Uses → anti hypertensive (Hyftonin)
① + C.HCl + KClO ₄ → Blue color	① Reserpine → Reserpine acid + Methanol + Trimethyl benzene ② Reserpamine → Reserpine acid + Methanol + Trimethyl benzene ③ Deserpine → Reserpine acid + Methanol + Trimethyl benzene

① Reserpine → Reserpine acid + Methanol + Trimethyl benzene
 ② Reserpamine → Reserpine acid + Methanol + Trimethyl benzene
 ③ Deserpine → Reserpine acid + Methanol + Trimethyl benzene

XII. Terpene alkaloids

Taxol
<chem>CN1[C@H]2CC[C@@H]3[C@H]4CC[C@@H]5[C@@]4(CC[C@H]3[C@H]2CC=C4[C@@]5(CC[C@@H](C4)O)C)C</chem>
in Taxus brevifolia (yew tree, death tree)
Uses → ① anticancer → TIL of ovarian cancer ② induce polymerization of protein tubulin to form stable non-F ₂ microtubules → stop cell division (mitosis) in cancer cell
Related cpd of pharmaceutical importance
① Taxotene → H ₂ O-soluble ② Taxotene → used as starting material for synthesis of Taxol